## INTERNATIONAL SEARCH REPORT

International application No. PCT/JP2004/014851

A.	CLASSIFICA	TION OF SUBJECT	T MATTER				
	Int.Cl7	A61K48/00,	31/7088,	35/76,	A61P35/00,	43/00,	35/02
		C12N15/09	·			•	•

According to International Patent Classification (IPC) or to both national classification and IPC

#### **B. FIELDS SEARCHED**

Minimum documentation searched (classification system followed by classification symbols)
Int.Cl<sup>7</sup> A61K48/00, 31/7088, 35/76, A61P35/00, 43/00, 35/02,
C12N15/09

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) CAPLUS (STN), MEDLINE (STN), BIOSIS (STN), EMBASE (STN)

### C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	ROSNET O. et al., Human FLT3/FLK2 Gene:cDNA Cloning and Expression in Hematopoietic Cells., Blood, 15 August, 1993 (15.08.93), 82(4), pages 1110 to 1119	1-13,16
Y	WO 2000/11470 Al (Chugai Pharmaceutical Co., Ltd.), 02 March, 2000 (02.03.00), Particularly, Claims; page 1, line 11 to page 3, line 10; page 9, line 25 to page 10, line 11 & EP 1109020 Al	1-13,16
Y	YEE Kevin W.H. et al., SU5416 and SU5614 inhibit kinase activity of wild-type and mutant FLT3 receptor tyrosine kinase., Blood, 15 October, 2002 (15.10.02), 100(8), p.2941-9	1-13,16

اث	Further documents are listed in the continuation of Box C.		See patent family annex.		
*A"	Special categories of cited documents: document defining the general state of the art which is not considered to be of particular relevance	«Ta	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention		
filing date  "L" document which may throw doubts on priority claim(s) or which is		"X"	considered novel or cannot be considered to involve an inventive step when the document is taken alone		
		"Y"			
"O" "P"	P" document published prior to the international filing date but later than the		combined with one or more other such documents, such combination being obvious to a person skilled in the art		
	priority date claimed	"&" document member of the same patent family			
	of the actual completion of the international search 04 January, 2005 (04.01.05)	Date	of mailing of the international search report 25 January, 2005 (25.01.05)		
Name and mailing address of the ISA/ Japanese Patent Office		Authorized officer			
Facsimile No.			Telephone No.		
orm F	CT/ISA/210 (second sheet) (January 2004)	<u> </u>			

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		P2004/014851			
C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT					
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.			
Y	YAMAMOTO Y. et al., Activating mutation of D835 within the activation loop of FLT3 in human hematologic malignancies., Blood, 15 April, 2001 (15.04.01), 97(8), p.2434-9	1-13,16			
Y	FIRE A. et al., Potent and specific genetic interference by double-stranded RNA in Caenorhabditis elegans., Nature, 19 February, 1998 (19:02.98), Vol.391, pages 806 to 811	1-13,16			
Y	DYKXHOORN D.M. et al., Killing the messenger: short RNAs that silence gene expression., Nature Rev.Mol.Cell Biol., 2003 June, Vol.4, pages 457 to 467	1-13,16			
Y	ELBASHIR S.M. et al., Duplexes of 21- nucleotide RNAs mediate RNA interference in cultured mammalian cells., Nature, 24 May, 2001 (24.05.01), Vol.411, pages 494 to 498	1-13,16			
Y	HEIDENREICH O. et al., AML1/MTG8 oncogene suppression by small interefering RNAs supports myeloid differentiation of t(8;21) - positive leukemic cells., Blood, 15 April, 2003 (15.04.03), 101(8), pages 3157 to 3163	1-13,16			
Y	MARTINEZ L.A. et al., Synthetic small inhibiting RNAs: Efficient tools to inactivate oncogenic mutations and restore p53 pathways., Proc.Natl.Acad.Sci.USA, 12 November, 2002 (12.11.02), Vol.99, pages 14849 to 14854	1-13,16			
Y	WILDA M. et al., Killing of leukemic cells with a BCR/ABL fusion gene by RNA interference (RNAi)., Oncogene, 2002, Vol.21, pages 5716 to 5724	1-13,16			
Y	SCHERR M. et al., Specific inhibition of bcr-abl gene expression by small interfering RNA., Blood, 15 February, 2003 (15.02.03), 101(4), pages 1566 to 1569	1-13,16			
Y	WOHLBOLD L. et al., Inhibition of bcr-abl gene expression by small interfering RNA sensitizes for imantinib mesylate(STI571)., Blood, 15 September, 2003 (15.09.03), 102(6), pages 2236 to 2239	1-13,16			
Y	SHEN C. et al., Gene Silencing by adenovirus-delivered siRNA., FEBS Letters, 27 March, 2002 (27.03.02), Vol.539, pages 111 to 114	1-13,16			
	10 (continuation of control 1 de la Virginia 2004)				

# INTERNATIONAL SEARCH REPORT

International application No. PCT/JP2004/014851

Box No. I	Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
1. X Claand this no Rule 2. \[ \]	national search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:  Claims Nos.: 14, 15  Decause they relate to subject matter not required to be searched by this Authority, namely:  ims 14 and 15 pertain to methods for treatment of the human body by therapy thus relate to a subject matter which this International Searching Authority of required, under the provisions of Article 17(2)(a)(i) of the PCT and 39.1(iv) of the Regulations under the PCT, to search.  Claims Nos.:  Decause they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box No. 1	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
1.	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.  As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.  As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
	No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark	The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

A. 発明の属する分野の分類 (国際特許分類 (IPC))

Int. Cl<sup>7</sup> A61K48/00, 31/7088, 35/76, A61P35/00, 43/00, 35/0<sup>2</sup>, C12N15/09

### B. 調査を行った分野

調査を行った最小限資料(国際特許分類(IPC))

Int. Cl<sup>7</sup> A61K48/00, 31/7088, 35/76, A61P35/00, 43/00, 35/0 2, C12N15/09

最小限資料以外の資料で調査を行った分野に含まれるもの

国際調査で使用した電子データベース (データベースの名称、調査に使用した用語)

CAPLUS (STN), MEDLINE (STN), BIOSIS (STN), EMBASE (STN)

### C. 関連すると認められる文献

引用文献の カテゴリー*	引用文献名 及び一部の箇所が関連するときは、その関連する箇所の表示	関連する 請求の範囲の番号
Y·	ROSNET O. et al., Human FLT3/FLK2 Gene:cDNA Cloning and Expression in Hematopoietic Cells., Blood, 1993 Aug 15, 82 (4), p.1110-1119	$1-13, \\ 16$
Y	WO 2000/11470 A1 (中外製薬株式会社),200 0.03.02,特に、特許請求の範囲、第1頁第11行〜第3頁 第10行及び第9頁第25行〜第10頁第11行 & EP 11 09020 A1	$1-13, \\ 16$
. Y	YEE Kevin W H et al., SU5416 and SU5614 inhibit kinase activ	1-13,

### 区欄の続きにも文献が列挙されている。

」 パテントファミリーに関する別紙を参照。.

- \* 引用文献のカテゴリー
- 「A」特に関連のある文献ではなく、一般的技術水準を示すもの
- 「E」国際出願日前の出願または特許であるが、国際出願日 以後に公表されたもの
- 「L」優先権主張に疑義を提起する文献又は他の文献の発行 日若しくは他の特別な理由を確立するために引用する 文献(理由を付す)
- 「O」口頭による開示、使用、展示等に言及する文献
- 「P」国際出願日前で、かつ優先権の主張の基礎となる出願

- の日の後に公表された文献
- 「T」国際出願日又は優先日後に公表された文献であって 出願と矛盾するものではなく、発明の原理又は理論 の理解のために引用するもの
- 「X」特に関連のある文献であって、当該文献のみで発明 の新規性又は進歩性がないと考えられるもの
- 「Y」特に関連のある文献であって、当該文献と他の1以上の文献との、当業者にとって自明である組合せによって進歩性がないと考えられるもの
- 「&」同一パテントファミリー文献

	関連すると認められる文献	
引用文献の   カテゴリー*	引用文献名 及び一部の箇所が関連するときは、その関連する箇所の表示	関連する 請求の範囲の番号
	ity of wild-type and mutant FLT3 receptor tyrosine kinase., Blood, 2002 Oct 15, 100 (8), p.2941-9	1 6
	YAMAMOTO Y et al., Activating mutation of D835 within the activation loop of FLT3 in human hematologic malignancies., Blood, 2001 Apr 15, 97 (8), p.2434-9	1-13, $16$
	FIRE A. et al., Potent and specific genetic interference by double-stranded RNA in Caenorhabditis elegans., Nature, 1998 Feb 19, Vol. 391, p. 806-811	$1-13, \\ 16$
	DYKXHOORN D. M. et al., Killing the messenger:short RNAs that silence gene wxpression., Nature Rev. Mol. Cell Biol., 2003 June, Vol. 4, p. 457-467	$1-13, \\ 16$
	ELBASHIR S. M. et al, Duplexes of 21-nucleotide RNAs mediate RNA interference in cultured mammalian cells., Nature, 2001 May 24, Vol. 411, p. 494-498	1-13, 16
·	HEIDENREICH O. et al., AML1/MTG8 oncogene suppression by small interefering RNAs supports myeloid differentiation of t(8;21)-positive leukemic cells., Blood, 2003 Apr 15, 101 (8), p. 3157-3163	$1-13, \\ 16$
	MARTINEZ L. A. et al., Synthetic small inhibiting RNAs: Efficient tools to inactivate oncogenic mutations and restore p53 pathways., Proc. Natl. Acad. Sci. USA, 2002 Nov 12, Vol. 99, p. 14849-14854	1-13, 16
	WILDA M. et al., Killing of leukemic cells with a <i>BCR/ABL</i> fusion gene by RNA interference(RNAi)., Oncogene, 2002, Vol. 21, p. 5716-5724	1-13, 16
	SCHERR M. et al., Specific inhibition of <i>bcr-abl</i> gene expression by small interfering RNA., Blood, 2003 Feb 15, 101 (4), p. 1566-1569	$1-13, \\ 16$
	WOHLBOLD L. et al., Inhibition of bcr-abl gene expression by small interfering RNA sensitizes for imantinib mesylate(STI571)., Blood, 2003 Sep 15, 102 (6), p. 2236-2239	$1-13, \\ 16$
l l	SHEN C. et al., Gene silencing by adenovirus-delivered siRNA., FEBS Letters, 2002 Mar 27, Vol. 539, p. 111-114	$1-13, \\ 16$
		-

第Ⅱ欄 請求の範囲の一部の調査ができないときの意見 (第1ページの2の続き)
法第8条第3項(PCT17条(2)(a))の規定により、この国際調査報告は次の理由により請求の範囲の一部について作成しなかった。
1. $oxed{oxed}$ 請求の範囲 $\phantom{aaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa$
請求の範囲14,15は、治療による人体の処置方法に関するものであって、PCT 17条(2)(a)(i)及びPCT規則39.1(iv)の規定により、この国際調 査機関が国際調査を行うことを要しない対象に係るものである。
2. □ 請求の範囲は、有意義な国際調査をすることができる程度まで所定の要件を満たしていない国際出願の部分に係るものである。つまり、
3. □ 請求の範囲は、従属請求の範囲であってPCT規則6.4(a)の第2文及び第3文の規定に 従って記載されていない。
第Ⅲ欄 発明の単一性が欠如しているときの意見(第1ページの3の続き)
次に述べるようにこの国際出願に二以上の発明があるとこの国際調査機関は認めた。
•
$\cdot$
1.   出願人が必要な追加調査手数料をすべて期間内に納付したので、この国際調査報告は、すべての調査可能な請求 の範囲について作成した。
2. □ 追加調査手数料を要求するまでもなく、すべての調査可能な請求の範囲について調査することができたので、追加調査手数料の納付を求めなかった。
3.   出願人が必要な追加調査手数料を一部のみしか期間内に納付しなかったので、この国際調査報告は、手数料の納付のあった次の請求の範囲のみについて作成した。
4.
追加調査手数料の異識の申立てに関する注意
□ 追加調査手数料の納付と共に出願人から異議申立てがあった。
追加調査手数料の納付と共に出願人から異議申立てがなかった。